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FIG. 1a(1)

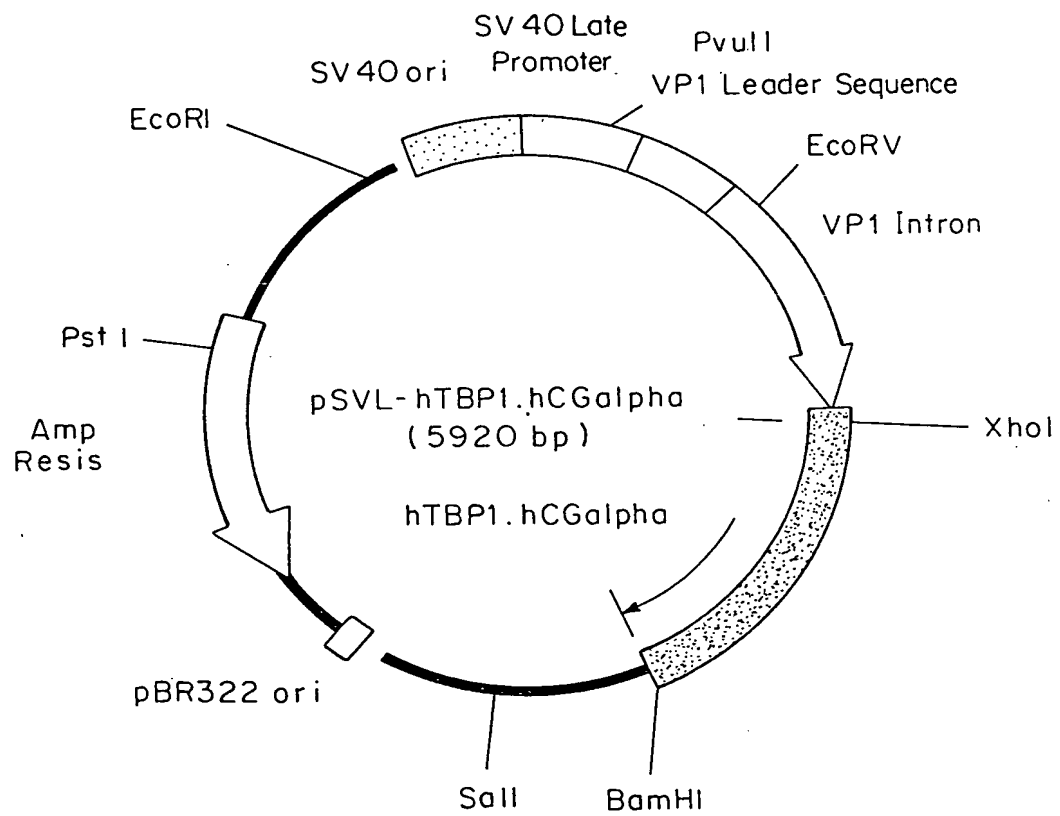


FIG. 10(2)

Xho I hGH Signal Sequence

hGH Intron

TCGAG ATG GCT ACA G GTAAGCGCCCTAAATCCCTTTGGGCACAATGTGTCTCTGAGGGGAGAGGCGACGACCTGTAGATGGACGGGGGCACCTAACCCCTCAGTTTGGGGTTTCT

Met Ala Thr

GAATGTAGTATCGCCATGTAAAGCCAGCATTTTGGCCCAATCTCAGAAAGCTCTGGTCTCTGGAGGATGGAGAGAGAAAACAAACAGCTCTGGAGCAGGAGAGTGTGGCCTCTCTGCTTTC

CGGCTCCCTCTGTGGCCCTCTGTGGTTTCTCCCCAGGC TCC CGG ACG TCC CTG CTC CTG GCT TTT GGC CTG TGC CTG CCC TGG CTT
Ser Arg Thr Ser Leu Leu Leu Ala Phe Gly Leu Leu Cys Leu pro Trp Leu

+20 Asp of Processed TBP1

CAA GAG GGC AGT GCC GAT AGT GTG TGT CCC CAA GGA AAA TAT ATC CAC CCT CAA AAT AAT TCG ATT TGC TGT ACC AAG TGC CAC AAA AAA
Gln Glu Gly Ser Ala Asp Ser Val Cys pro Gln Gly Lys Tyr Ile His pro Gln Asn Ser Ile Cys Cys Thr Lys Cys His Lys Gly

ACC TAC TTG TAC AAT GAC TGT CCA GGC CCG GGC CAG GAT ACG GAC TGC AGG GAG TGT GAG AGC GGC TCC TTC ACC GCT TCA GAA AAC CAC CTC
Thr Tyr Leu Tyr Asn Asp Cys pro Gly pro Gly Gln Asp Thr Asp Cys Arg Glu Cys Glu Ser Gly Ser Phe Thr Ala Ser Glu Asn His Leu

AGA CAC TGC CTC AGC TGC TCC AAA TGC CGA AAG GAA ATG GGT CAG GTG GAG ATC TCT TGC ACA GTG GAC CGG GAC ACC GTG TGT GGC TGC
Arg His Cys Leu Ser Cys Ser Lys Cys Arg Lys Glu Met Gly Gln Val Glu Ile Ser Ser Cys Thr Val Asp Arg Asp Thr Val Cys Gly Cys

AGG AAG AAC CAG TAC CGG CAT TAT TGG AGT GAA AAC CTT TTC CAG TGC TTC AAT TGC AGC CTC TGC AAT GGG ACC GTG CAC CTC TCC TGT
Arg Lys Asn Gln Tyr Arg His Tyr Trp Ser Glu Asn Leu Phe Gln Cys Phe Asn Cys Ser Leu Cys Leu Asn Gly Thr Val His Leu Ser Cys

Linker

CAG GAG AAA CAG AAC ACC GTG TGC ACC TGC CAT GCA GGT TTC TTT CTA AGA GAA AAC GAG TGT GTC TCC TGT GCC GGT GCT GCC CCA GGT
Gln Glu Lys Gln Asn Thr Val Cys Thr Cys His Ala Gly Phe Leu Arg Glu Asn Glu Cys Val Ser Cys Ala Gly Ala Ala Pro Gly

+7 Cys of hCG alpha

TGC CCA GAA TGC ACG CTA CAG GAA AAC CCA TTC TCC CAG CCG GGT GGC CCA ATA CTT CAG TGC ATG GGC TGC TGC TTC TCT AGA GCA TAT
Cys pro Glu Cys Thr Leu Gln Glu Asn pro Phe Ser Gln pro Gly Ala pro Ile Leu Gln Cys Met Gly Cys Phe Ser Arg Ala Tyr

CCC ACT CCA CTA AGG TCC AAG AAG ACG ATG TTG GTC CAA AAG AAC GTC ACT TCA GAG TCC ACT TGC TGT GTA GCT AAA TCA TAT AAC AGG GTC
Pro Thr pro Leu Arg Ser Lys Lys Thr Met Leu Val Gln Lys Asn Val Thr Ser Glu Ser Thr Cys Cys Val Ala Lys Ser Tyr Asn Arg Val

ACA GTA ATG GGG GGT TTC AAA GTG GAG AAC CAC ACG GCG TGC CAC TGC AGT ACT TGT TAT TAT CAC AAA TCT TAA G

Thr Val Met Gly Gly Phe Lys Val Glu Asn His Thr Ala Cys His Cys Ser Thr Cys Tyr Tyr His Lys Ser ... |

Bam HI

FIG. 1b(1)

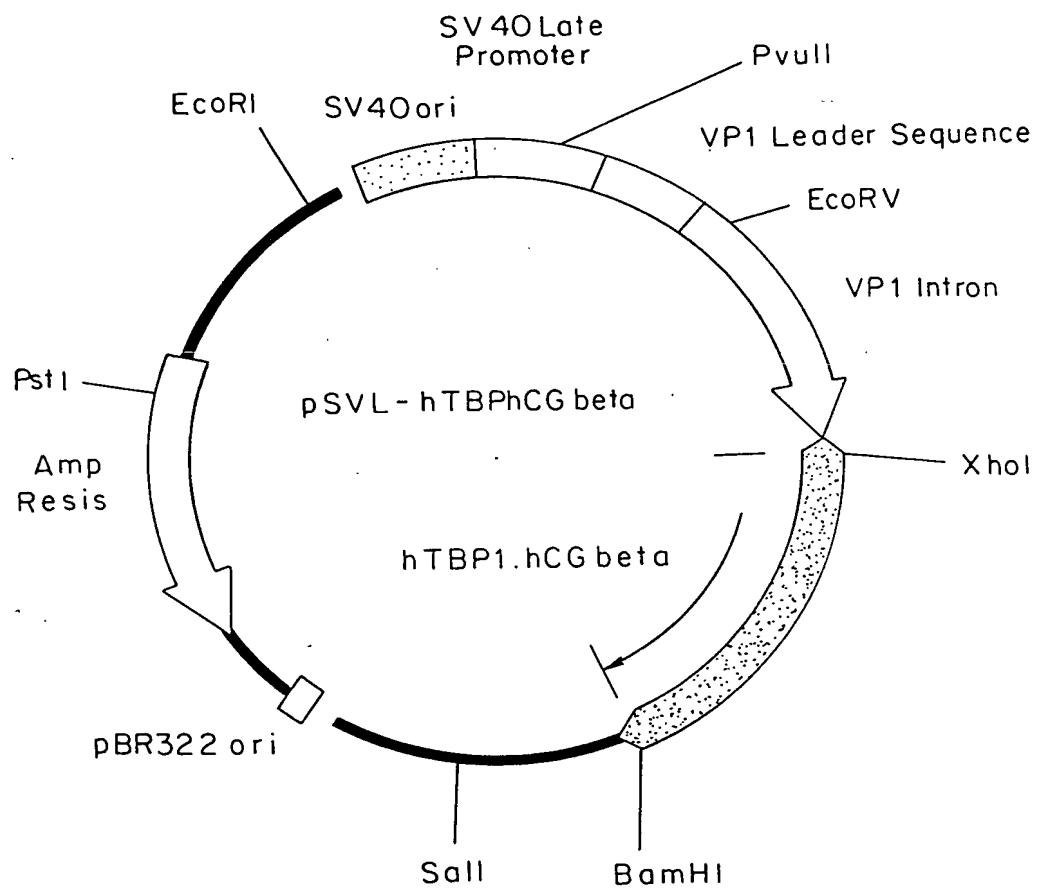


FIG. 1b(2)

XhoI
 CTCGAG ATG GCT ACA G GTRAGCGCCCTAAATCCCTTTGGGCACAATGTCTCTAGGGGAGAGTAGGACGACCTGTAGTGGGACGGGGGCACTAACCCCTCAGGTTTGGG
 ▶ Met, Ala Thr

hGH Signal Sequence
 hGH Intron

GCTTCTGAATGTGAGTATGCGCCATGTATAGCCCGATTTTGGCCCAATCTCAGAAAGCTCCTGGTCCCTGGAGGATGGAGAGAGAAAACAAACAGCTCCTGGAGCAGGGAGAGTGTCTGAC
 CTCTGTCTCCGGCTCCCTCTGTGTGCTCTGTGCTCTGTGTTCTCTCCCGAGGC TCC CGG ACG TCC CTG CTG GCT TTT GGC CTG CTC TGC CTG
 ▶ Ser Arg Thr Ser Leu Leu Ala Phe Gly Leu Leu Cys Leu

+20 Asp of Processed TBPI

CCC TGG CTT CAA GAG GGC AGT GCC GAT AGT GTG TGT CCC CAA GGA AAA TAT ATC CAC CCT CAA AAT AAT TCG ATT TGC TGT ACC
 ▶ Pro Trp Leu Gln Glu Gly Ser Ala Asp Ser Val Cys Pro Gln Gly Tyr Ile His Pro Gln Asn Ser Ile Cys Cys Thr

AAG TGC CAC AAA GGA ACC TAC TTG TAC AAT GAC TGT CCA GGC CCG GGC CAG GAT ACG GAC TGC AGG GAG TGT GAG AGC GGC TCC TTC ACC
 ▶ Lys Cys His Lys Gly Thr Tyr Leu Tyr Asn Asp Cys Pro Gly Pro Gly Gln Asp Thr Asp Cys Arg Glu Cys Glu Ser Gly Ser Phe Thr

GCT TCA GAA AAC CAC CTC AGA CAC TGC CTC AGC TGC TCC AAA TGC CGA AAG AAG ATG GGT CAG GTG GAG ATC TCT TCT TGC ACA GTG GAC
 ▶ Ala Ser Glu Asn His Leu Arg His Cys Leu Ser Cys Ser Lys Cys Arg Lys Glu Met Gly Gln Val Ile Ser Ser Cys Thr Val Asp

CGG GAC ACC GTG TGT GGC TGC AGG AAG AAC CAG TAC CGG CAT TAT TGG AGT GAA AAC CTT TTC CAG TGC TTC AAT TGC AGC CTC TGC CTC
 ▶ Arg Asp Thr Val Cys Gly Cys Arg Lys Asn Gln Tyr Arg His Tyr Trp Ser Glu Asn Leu Phe Gln Cys Phe Asn Cys Ser Leu Cys Leu

AAT GGG ACC GTG CAC CTC TCC TGC CAG GAG AAA CAG AAC ACC GTG TGC ACC TGC CAT GCA GGT TTC TTT CTA AGA GAA AAT GAG TGT GTC
 ▶ Asn Gly Thr Val His Leu Ser Cys Gln Glu Lys Gln Asn Thr Val Cys Thr Cys His Ala Gly Phe Leu Arg Glu Asn Glu Cys Val

Linker
 +7 Pro of hCG beta

TCC TGT GCT GGT GCT CCA CGG TGC CGC CCC ATC AAT GCC ACC CTG GCT GTG GAG AAG GAG GGC TGC CCC GTG TGC ATC ACC GTC
 ▶ Ser Cys Ala Gly Ala Gly Pro Arg Cys Arg Pro Ile Asn Ala Thr Leu Ala Val Glu Lys Glu Gly Cys Pro Val Cys Ile Thr Val

AAC ACC AAC ATC TGT GCC GGC TAC TGC CCC ACC ATG ACC CGC GTG CAG GGG GTC CTG CCG GCC CTG CCT CAG GTG GTG TGC AAC TAC
 ▶ Asn Thr Thr Ile Cys Ala Gly Tyr Cys Pro Thr Met Thr Arg Val Leu Gln Gly Val Leu Pro Ala Leu Pro Gln Val Val Cys Asn Tyr

CGC GAT GTG CGC TTC GAG TCC ATC CGG CTC CCT GGC TGC CCG CGC GTG AAC CCC GTG GTC TCC TAC GCC GTG GCT CTC AGC TGT CAA
 ▶ Arg Asp Val Arg Phe Glu Ser Ile Arg Leu Pro Gly Cys Pro Arg Gly Val Val Ser Tyr Ala Val Ala Leu Ser Cys Gln

TGT GCA CTC TGC CGC CGC AGC ACC ACT GAC TGC GGC GGT CCC AAG GAC CAC CCC TTG ACC TGT GAT GAC CCC CGC TTC CAG GAC TCC TCT
 ▶ Cys Ala Leu Cys Arg Arg Ser Thr Thr Asp Cys Gly Gly Pro Lys Asp His Pro Leu Thr Cys Asp Pro Arg Phe Gln Asp Ser Ser

TCC TCA AAG GCC CCT CCC AGC CTT CCA AGC CCA TCC CGA CTC CCG GCG CCC TCG GAC ACC CCG ATC CTC CCA CAA TAA
 ▶ Ser Ser Lys Ala Pro Pro Ser Leu Pro Ser Pro Ser Arg Leu Pro Gly Pro Ser Asp Thr Pro Ile Leu Pro Gln ***

FIG. 2a(1)

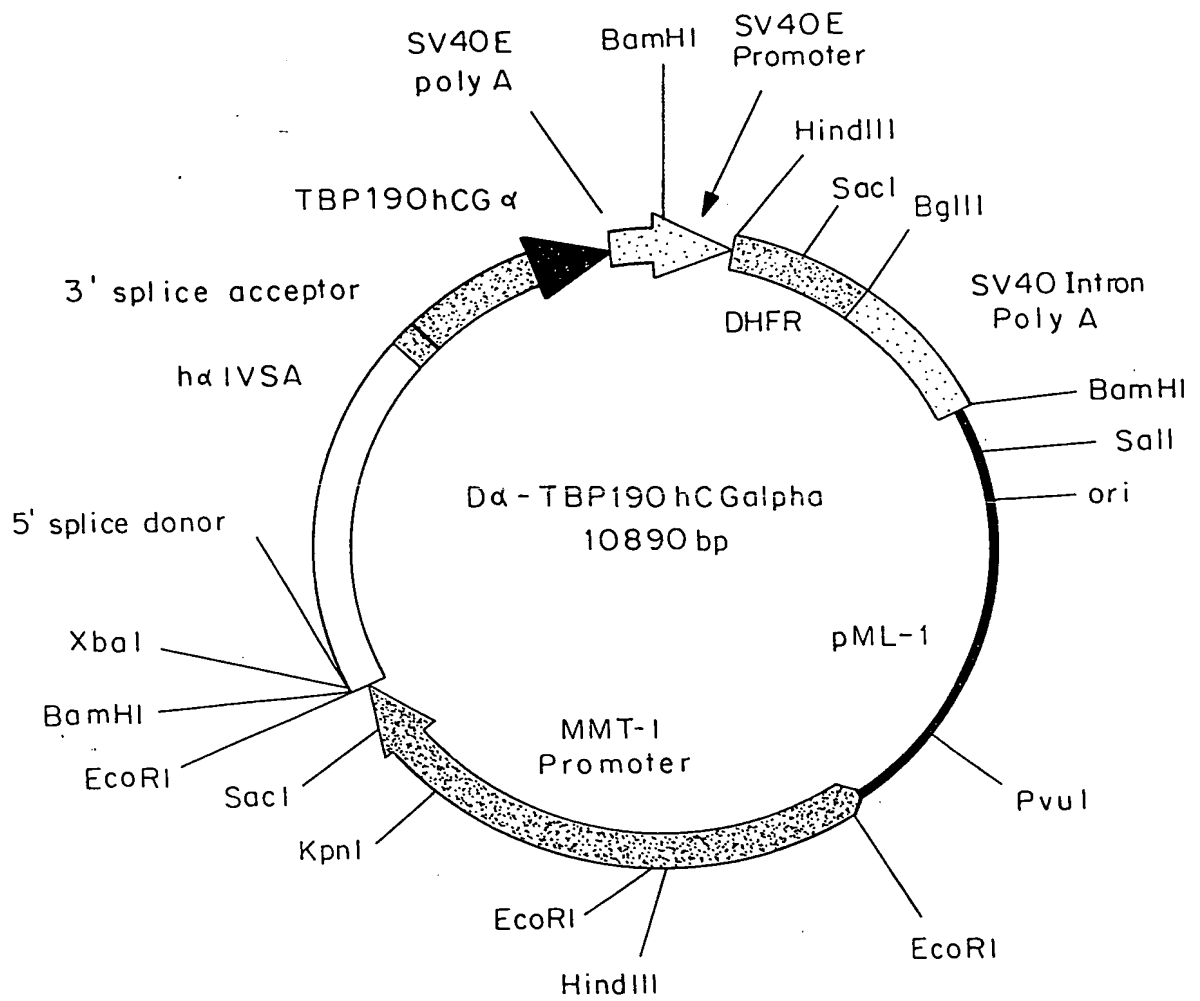


FIG. 2a(2)

XhoI hGH Signal Sequence hGH Intron

TCGAG ATG GCT ACA G GTAAGCGCCCTAAATCCCTTTGGGACAAATGTCCTGAGGGAGAGGACGACCTGTAGTGGACCGGGGGGCACTAACCTCAGGTTTGGGGTTTCT

► Met Ala Thr

GAATGTGATCGCCATGTAAGCCCGCAGTATTGGCCAATCTCAGAAAGCTCCTGGTCCCTGGAGGGATGGAGAGAAACAAACAGCTCCTGGACAGGAGAGTGCTGGCCTCTTGCTCTC

CGGCTCCCTCTGTGGCCCTCTGGTTCTCCCGAGGC TCC CGG ACG TCC CTG CTC CTG GCT TTT GGC CTG CTC TGC CTG CCC TGG CTT

► Ser Arg Thr Ser Leu Leu Ala Phe Gly Leu Cys Leu Pro Trp Leu

+20 Asp of processed TBPI

CAA GAG GGC AGT GCC GAT AGT GTG TGT CCC CAA GGA AAA TAT ATC CAC CCT CAA AAT AAT TCG ATT TGC TGT ACC AAG TGC CAC AAA GGA

► Gln Glu Gly Ser Ala Asp Ser Val Cys Pro Gln Gly Lys Tyr Ile His pro Gln Asn Ser Ile Cys Cys Thr Lys Cys His Lys Gly

ACC TAC TTG TAC AAT GAC TGT CCA GGC CCG GGC CAG GAT ACG GAC TGC AGG GAG TGT GAG AGC GGC TCC TTC ACC GCT TCA GAA AAC CAC CTC

► Thr Tyr Leu Tyr Asn Asp Cys Pro Gly Pro Gly Gln Asp Thr Asp Cys Arg Glu Cys Glu Ser Gly Ser Phe Thr Ala Ser Glu Asn His Leu

AGA CAC TGC CTC AGC TGC TCC AAA TGC CGA AAG GAA ATG GGT CAG GTG GAG ATC TCT TCT TGC ACA GTG GAC CGG GAC ACC GTG TGT GGC TGC

► Arg His Cys Leu Ser Cys Ser Lys Cys Arg Lys Glu Met Gly Gln Val Glu Ile Ser Ser Cys Thr Val Asp Arg Asp Thr Val Cys Gly Cys

AGG AAG AAC CAG TAC CGG CAT TAT TGG AGT GAA AAC CTT TTC CAG TGC TGC AAT TGC AGC CTC TGC CTC AAT GGG ACC GTG CAC CTC TCC TGC

► Arg Lys Asn Gln Tyr Arg His Tyr Trp Ser Glu Asn Leu Phe Gln Cys Phe Asn Cys Ser Leu Cys Leu Asn Gly Thr Val His Leu Ser Cys

CAG GAG AAA CAG AAC ACC GTG TGC ACC TGC CAT GCA GGT TTC TTT CTA AGA GAA AAC GAG TGT GTC TCC TGT AGT AAC TGT AAG AAA AGC CTG

► Gln Glu Lys Gln Asn Thr Val Cys Thr Cys His Ala Gly Phe Leu Arg Glu Asn Glu Cys Val Ser Cys Ser Asn Cys Lys Lys Ser Leu

Linker +7 Cys of hCG alpha

GAG TGC ACG AAG TTG TGC CTA CCC CAG ATT GAG AAT GTT AAG GGC ACT GAG GAC TCA GGC ACC ACA GCC GGT GCT GCC CCA GGT TGC CCA

► Glu Cys Thr Lys Leu Cys Leu Pro Gln Ile Glu Asn Val Lys Gly Thr Glu Asp Ser Gly Thr Thr Ala Gly Ala pro Gly Cys Pro

GAA TGC ACG CTA CAG GAA AAC CCA TTC TCC CAG CCG GGT GCC CCA ATA CTT CAG TGC ATG GGC TGC TGC TTC TCT TCT AGA GCA TAT CCC ACT

► Glu Cys Thr Leu Gln Glu Asn Pro Phe Ser Gln Pro Gly Ala Pro Ile Leu Gln Cys Met Gly Cys Cys Phe Ser Arg Ala Tyr Pro Thr

CCA CTA AGG TCC AAG AAG ACG ATG TTG GTC CAA AAG AAC GTC ACC TCA GAG TCC ACT TGC TGT GTA GCT AAA TCA TAT AAC AGG GTC ACA GTA

► Pro Leu Arg Ser Lys Lys Thr Met Leu Val Gln Lys Asn Val Thr Ser Glu Ser Thr Cys Cys Val Ala Lys Ser Tyr Asn Arg Val Thr Val

ATG GGG GGT TTC AAA GTG GAG AAC CAC ACG GCG TGC CAC TGC AGT ACT TGT TAT CAC AAA TCT TAA GGATCCCTCGAG

► Met Gly Gly Phe Lys Val Glu Asn His Thr Ala Cys His Cys Ser Thr Cys Tyr Tyr His Lys Ser ***

Bam HI XhoI

FIG. 2b(1)

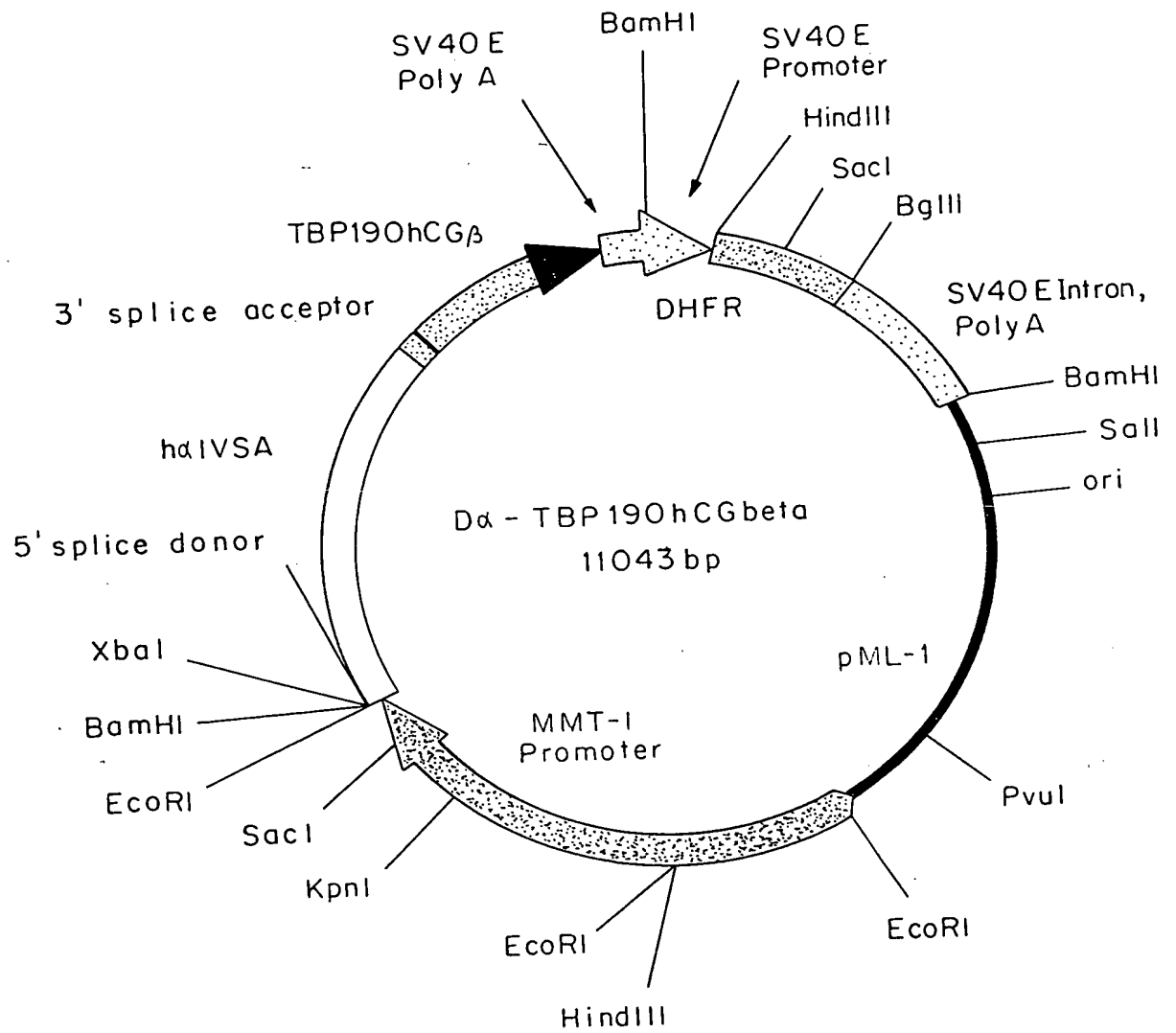


FIG. 2b(2)

XhoI hGH-Signal Sequence hGH Intron
CTCGAG ATG GCT ACA G GTAAGCGCCCTAAATCCCTTTGGGCACAATGTCTCCTGAGGGGAGACGACGACCTGTAGTGGACGGGGGCACCTAACCTCAGGTTTGGG
▶ Met Ala Thr
GCTTCTGATGTGATGATCGCCATCGCCATGTAAGCCAGTATTGGCCCATCTCAGAAAGCTCTGGTCCCTGGAGGATGGAGAGAGAAAAACAACAGCTCCTGTAGCAGGTAGAGTCTCTGGC
CTCTTGCTCTCCGGCTCCCTCTGTGTCCTCTGTTCCCTCTGTTCTTCTCCCCAGG C TCC CGG AGG TCC CTG CTG GCT TTT GGC CTG CTG CTG CTG
▶ Ser Arg Thr
+20 Asp of Processed TBP1
CCC TGG CTT CAA GAG GGC AGT GCC GAT AGT GTG TGT CCC CAA GGA AAA TAT ATC CAC CCT CAA AAT AAT TCG ATT TGC TGT ACC
▶ Pro Trp Leu Gln Glu Gly Ser Ala Asp Ser Val Cys Pro Gln Gly Lys Tyr Ile His Pro Gln Asn Ser Ile Cys Cys Thr
AAG TGC CAC AAA GGA ACC TAC TTG TAC AAT GAC TGT CCA GGC CCG GGC CAG GAT ACG GAC TGC AGG GAG TGT GAG AGC GGC TCC TTC ACC
▶ Lys Cys His Lys Gly Thr Tyr Leu Tyr Asn Asp Cys Pro Gly Pro Gly Gln Asp Thr Asp Cys Arg Glu Cys Glu Ser Gly Ser Phe Thr
GCT TCA GAA AAC CAC CTC AGA CAC TGC CTC AGC TGC TCC AAA TGC CGA AAG GAA ATG GGT CAG GTG GAG ATC TCT TCT TGC ACA GTG GAC
▶ Ala Ser Glu Asn His Leu Arg His Cys Leu Ser Cys Ser Lys Cys Arg Lys Glu Met Gly Gln Val Glu Ile Ser Ser Cys Thr Val Asp
CGG GAC ACC GTG TGT GGC TGC AGG AAG AAC CAG TAC CGG CAT TAT TGG AGT GAA AAC CTT TTC CAG TGC TTC AAT TGC AGC CTC TGC CTC
▶ Arg Asp Thr Val Cys Gly Cys Arg Lys Asn Gln Tyr Arg His Tyr Trp Ser Glu Asn Leu Phe Gln Cys Phe Asn Cys Ser Leu Cys Leu
AAT GGG ACC GTG CAC CTC TCC TGC CAG GAG AAA CAG AAC ACC GTG TGC ACC CAT CCA GGT TTC TTT CTA AGA GAA AAC GAG TGT GTC
▶ Asn Gly Thr Val His Leu Ser Cys Gln Glu Lys Gln Asn Thr Val Cys Thr Cys His Ala Gly Phe Leu Arg Glu Asn Glu Cys Val
TCC TGT AGT AAC TGT AAG AAA AGC CTG GAG TGC ACG AAG TTG TGC CTA CCC CAG ATT GAG AAT GTT AAG GGC ACT GAG GAC TCA GGC ACC
▶ Ser Cys Ser Asn Cys Lys Lys Ser Leu Glu Cys Thr Lys Leu Cys Leu Pro Gln Ile Glu Asn Val Lys Gly Thr Glu Asp Ser Gly Thr
Linker
+7 Pro of beta
ACA GCT GGT GCT GGT CCA CGG TGC CGC CCC ATC AAT GCC ACC CTG GCT GTG GAG AAG GGC TGC CCC CTG TGC ATC ACC GTC AAC
▶ Thr Ala Gly Ala Gly Pro Arg Cys Arg Pro Ile Asn Ala Thr Leu Ala Val Glu Lys Glu Gly Cys Pro Val Cys Ile Thr Val Asn
ACC ACC ATC TGT GCC GGC TAC TGC CCC ACC ATG ACC CGC GTG CTG CAG GGC GTG CTC CCG GCC CTG CCT CAG GTG TGC AAC TAC CGC
▶ Thr Thr Ile Cys Ala Gly Tyr Cys Pro Thr Met Thr Arg Val Leu Gln Gly Val Leu Pro Ala Leu Pro Gln Val Val Cys Asn Tyr Arg
GAT GTG CGC TTC GAG TCC ATC CGG CTC CCT GGC TGC CCG GGC GTG AAC CCC GTG GTC TCC TAC GCC GTG GCT CTC AGC TGT CAA TGT
▶ Asp Val Arg Phe Glu Ser Ile Arg Leu Pro Gly Cys Pro Arg Gly Val Asn Pro Val Val Ser Tyr Ala Val Ala Leu Ser Cys Gln Cys
GCA CTC TGC CGC AGC ACC ACT GAC TGC GGC GGT CCC AAG GAC CAC CCC TTG ACC TGT GAT GAC CCC CGC TTC CAG GAC TCC TCT TCC
▶ Ala Leu Cys Arg Arg Ser Thr Thr Asp Cys Gly Gly Pro Lys Asp His Pro Leu Thr Cys Asp Asp Pro Arg Phe Gln Asp Ser Ser
TCA AAG GCC CCT CCC AGC CTT CCA AGC CCA TCC CGA CTC CCG GGC CCC TCG GAC ACC CCG ATC CTC CCA TAA GGATCCCTCGAG
▶ Ser Lys Ala Pro Pro Ser Leu Pro Ser Pro Ser Arg Leu Pro Gly Pro Ser Asp Thr Pro Ile Leu Pro Gln *** BamIII XhoI

FIG. 3

p55 TNFR1, TBP1 and TBP1 FUSION CONSTRUCTS

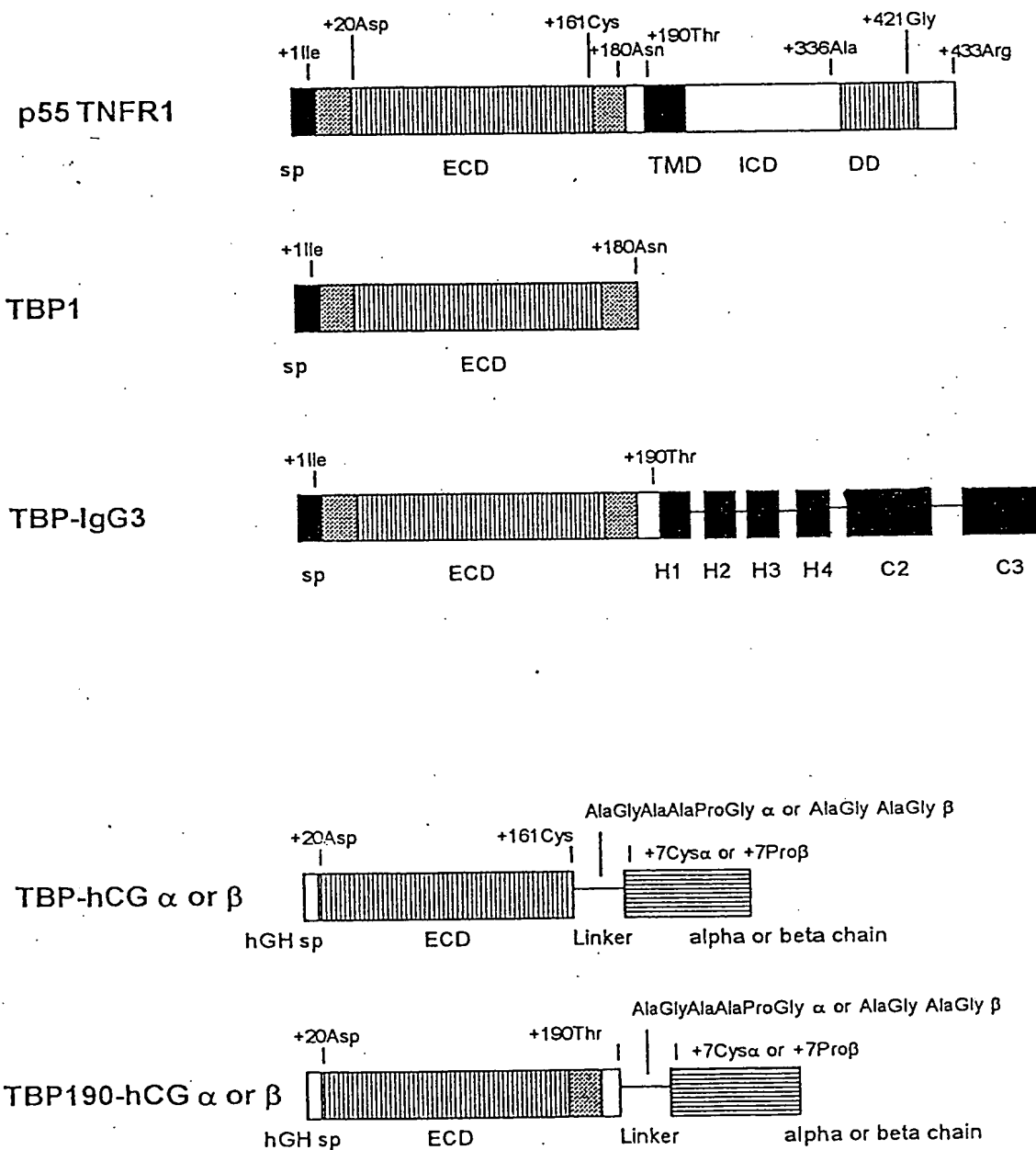
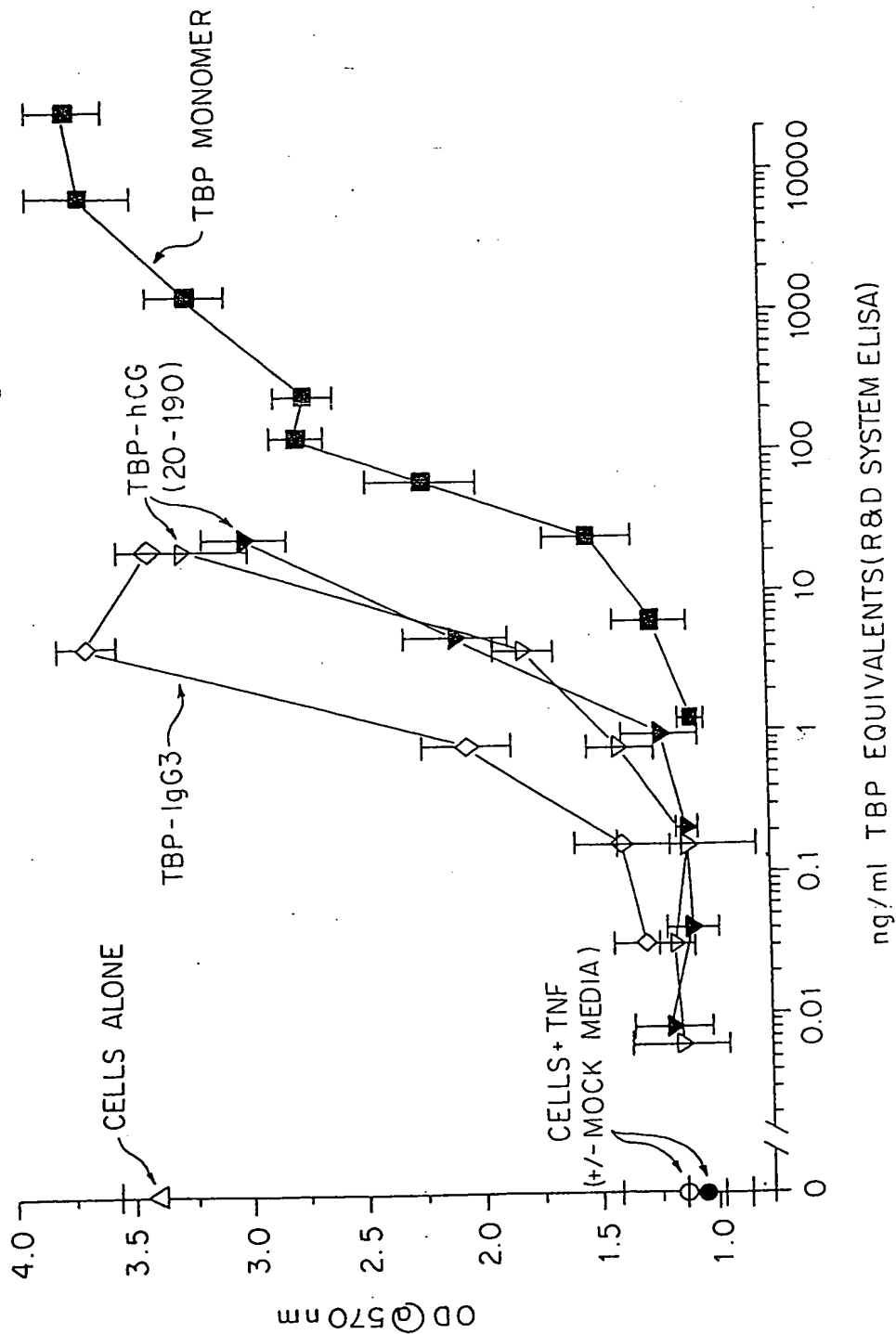
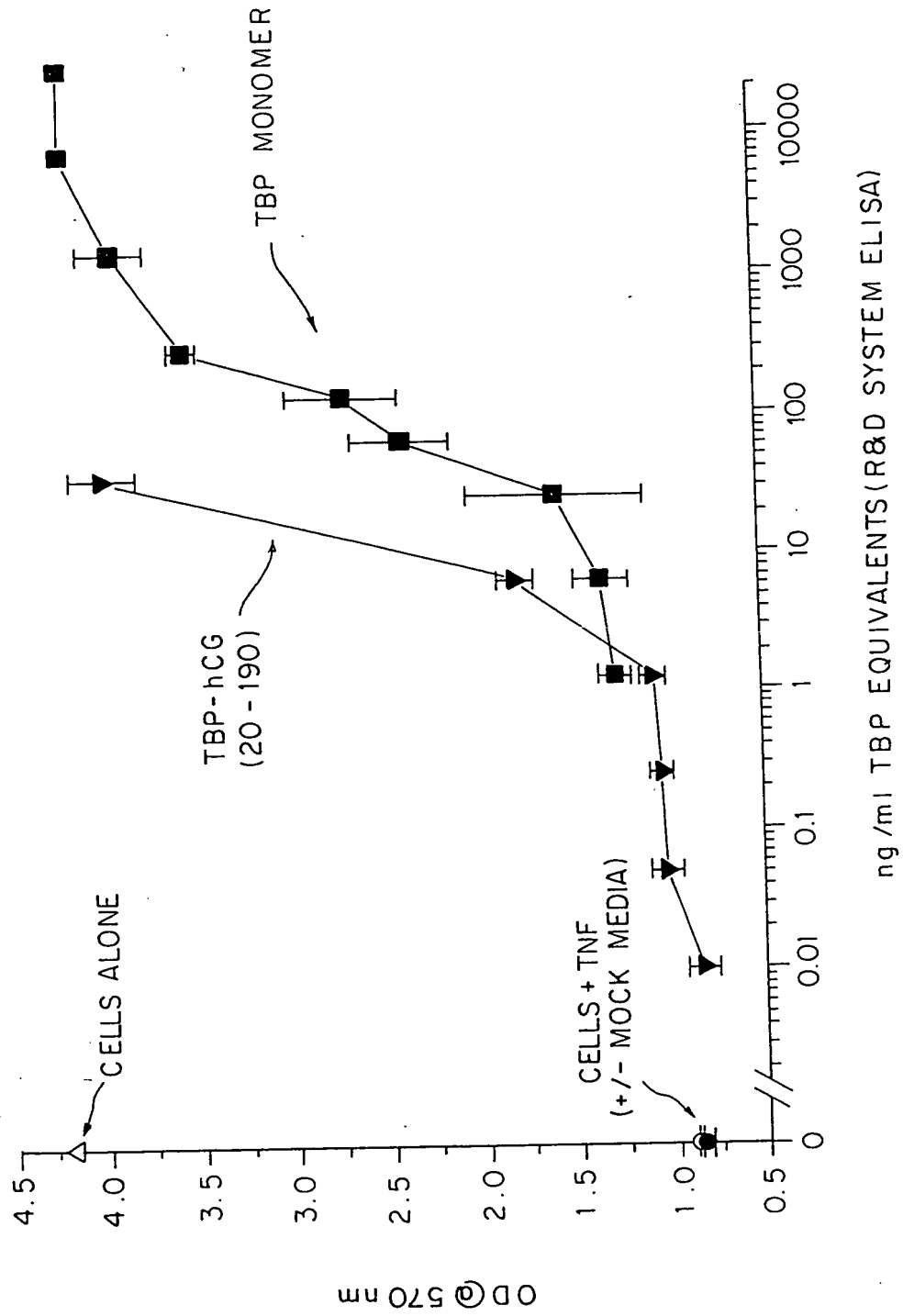


FIG. 4

- 10⁵ CELLS/WELL + 2.5 ng/ml TNF α + TBP MONOMER
- △ CELLS ALONE
- 2.5 ng/ml TNF α (NO TBP)
- ◇— TBP-IgG3 CHO MED + 2.5 ng/ml TNF α
- ▼— TBP-hCG (20-190) CHO MEDIA + 2.5 ng/ml TNF α
- ▽— TBP-hCG (20-190) CHO MEDIA + 2.5 ng/ml TNF α
- CELLS + CHO MOCK MEDIA + 2.5 ng/ml TNF α



- 10⁵ CELLS / WELL + 2.5 ng/ml TNF α + TBP MONOMER
 Δ CELLS ALONE
 ○ CELLS + 2.5 ng/ml TNF α (NO TBP)
 —▼— CELLS + TBP-hCG (20-190) COS7 MED + 2.5 ng/ml TNF α
 ● CELLS + COS7 MOCK TRANSFECTANT MEDIA + 2.5 ng/ml TNF α



- 10⁵ CELLS / WELL + 2.5 ng/ml TNF α + TBP MONOMER
 Δ CELLS ALONE
 ○ CELLS + 2.5 ng/ml TNF α (NO TBP)
 —▽— CELLS + PURIFIED TBP-hCG (20-161)

FIG. 6

